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| <b>TRANSMITTAL LETTER TO THE UNITED STATES<br/>DESIGNATED/ELECTED OFFICE (DO/EO/US)<br/>CONCERNING A SUBMISSION UNDER 35 U.S.C. 371</b>  |   | ATTORNEY'S DOCKET NUMBER<br><b>252412</b>                                     |
|  |   | CLIENT REF. NO.<br><b>PA36014US-FZJH903ver</b>                                |
|  |   | U.S. APPLICATION NO. (if known see 37 CFR 1.5)<br><b>Unassigned 10/580992</b> |
| INTERNATIONAL APPLICATION NO.<br><b>PCT/EP2004/013451</b>  | INTERNATIONAL FILING DATE<br><b>26 November 2004 (26.11.2004)</b> | PRIORITY DATE CLAIMED<br><b>26 November 2003 (26.11.2003)</b>                 |
| TITLE OF INVENTION<br><b>NOVEL MACROCYCLES FOR THE TREATMENT OF CANCER</b>   |   |   |
| APPLICANT(S) FOR DO/EO/US<br><b>WESSJOHANN et al.</b>  |   |   |
| Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:  |   |   |
| 1. <input checked="" type="checkbox"/> This is a <b>FIRST</b> submission of items concerning a filing under 35 USC 371 and 37 CFR 1.491.   |   |   |
| 2. <input type="checkbox"/> This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a filing under 35 USC 371 and 37 CFR 1.491.  |   |   |
| 3. <input type="checkbox"/> This is an express request to begin national examination procedures (35 USC 371(f)).   |   |   |
| 4. <input type="checkbox"/> The US has been elected (PCT Article 31).  |   |   |
| 5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 USC 371(c)(2)) <ul style="list-style-type: none"> <li>a. <input type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input checked="" type="checkbox"/> has been communicated by the International Bureau.</li> <li>c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</li> </ul>  |   |   |
| 6. <input checked="" type="checkbox"/> An English language translation of the International Application as filed (35 USC 371(c)(2)). <ul style="list-style-type: none"> <li>a. <input checked="" type="checkbox"/> is attached hereto.</li> <li>b. <input type="checkbox"/> has previously been submitted under 35 U.S.C. 154(d)(4).</li> </ul>  |   |   |
| 7. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 USC 371(c)(3)) <ul style="list-style-type: none"> <li>a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input type="checkbox"/> have been communicated by the International Bureau.</li> <li>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</li> <li>d. <input checked="" type="checkbox"/> have not been made and will not be made.</li> </ul> |   |   |
| 8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 USC 371(c)(3)).   |   |   |
| 9. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 USC 371(c)(4)).  |   |   |
| 10. <input type="checkbox"/> An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 USC 371(c)(5)).   |   |   |
| 11. Nucleotide and/or Amino Acid Sequence Submission <ul style="list-style-type: none"> <li>a. <input type="checkbox"/> Computer Readable Form (CRF)</li> <li>b. Specification Sequence Listing on:             <ul style="list-style-type: none"> <li>i. <input type="checkbox"/> CD-ROM or CD-R (2 copies); or</li> <li>ii. <input type="checkbox"/> Paper Copy</li> </ul> </li> <li>c. <input type="checkbox"/> Statement verifying identity of above copies</li> </ul>   |   |   |
| <b>Items 12 to 20 below concern other document(s) or information included:</b>   |   |   |
| 12. <input checked="" type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98. <ul style="list-style-type: none"> <li><input checked="" type="checkbox"/> Form PTO-1449</li> <li><input checked="" type="checkbox"/> Copies of References (except for U.S. patents and applications)</li> </ul>   |   |   |
| 13. <input type="checkbox"/> An assignment for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.  |   |   |
| 14. <input checked="" type="checkbox"/> A <b>FIRST</b> preliminary amendment.  |   |   |
| 15. <input type="checkbox"/> A <b>SECOND</b> or <b>SUBSEQUENT</b> preliminary amendment.   |   |   |
| 16. <input type="checkbox"/> A substitute specification.   |   |   |
| 17. <input type="checkbox"/> A change of power of attorney and/or address letter.  |   |   |
| 18. <input checked="" type="checkbox"/> Application Data Sheet Under 37 CFR 1.76   |   |   |
| 19. <input checked="" type="checkbox"/> Return Receipt Postcard  |   |   |
| 20. <input type="checkbox"/> Other items or information:   |   |   |

**SEND TO: Mail Stop PCT, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450**

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| U.S. APPLICATION NO. (if known, see 37 CFR 1.5)<br><b>Unassigned</b><br><b>10/580992</b> |  | INTERNATIONAL APPLICATION NO.<br>PCT/EP2004/013451 |  | ATTORNEY DOCKET NO.<br>252412 |  | CLIENT REF. NO.<br>PA36014US-<br>FZJH903ver |  |
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|  |              |   |         |                       |              |
|--|--------------|---|---------|-----------------------|--------------|
| The following fees are submitted:  |              |   |         | CALCULATIONS          | PTO USE ONLY |
| 21. <input checked="" type="checkbox"/> Basic national fee ..... \$300   |              |   |         | \$300.00              |              |
| 22. <input checked="" type="checkbox"/> Examination Fee  |              |   |         |                       |              |
| If international preliminary examination report prepared by USPTO and all claims satisfy provisions of PCT Article 33(1)-(4) ..... \$ 0  |              |   |         | \$200.00              |              |
| All other situations ..... \$200   |              |   |         |                       |              |
| 23. <input checked="" type="checkbox"/> Search Fee   |              |   |         |                       |              |
| USPTO was ISA ..... \$100  |              |   |         | \$500.00              |              |
| Search report provided to USPTO by other entity ..... \$400  |              |   |         |                       |              |
| All other situations ..... \$500   |              |   |         |                       |              |
| <b>TOTAL OF 21, 22 AND 23 =</b>  |              |   |         | <b>\$1,000.00</b>     |              |
| <input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. |              |   |         | \$                    |              |
| Total Sheets   | Extra Sheets | Number of each additional 50 or fraction thereof (round up to a whole number) | RATE    |                       |              |
| - 100 =  | / 50 =       |   | x \$250 | \$                    |              |
| <input type="checkbox"/> Surcharge of \$130 for furnishing the National fee or oath or declaration later than 30 months from the earliest claimed priority date (37 CFR 1.492(h)).   |              |   |         | \$                    |              |
| CLAIMS   | NUMBER FILED | NUMBER EXTRA  | RATE    |                       |              |
| Total Claims   | 13 - 20 =    | 0   | x \$ 50 | \$0.00                |              |
| Independent Claims   | 1 - 3 =      | 0   | x \$200 | \$0.00                |              |
| MULTIPLE DEPENDENT CLAIM(S) (if applicable)  |              |   |         | + \$360               | \$0.00       |
| <b>TOTAL OF ABOVE CALCULATIONS =</b>   |              |   |         | <b>\$1,000.00</b>     |              |
| <input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2.   |              |   |         | \$ ( )                |              |
| <b>SUBTOTAL=</b>   |              |   |         | <b>\$1,000.00</b>     |              |
| <input type="checkbox"/> Processing fee of \$130 for furnishing English Translation later than 30 months from the earliest claimed priority date.  |              |   |         | \$                    |              |
| <b>TOTAL NATIONAL FEE=</b>   |              |   |         | <b>\$</b>             |              |
| <input type="checkbox"/> Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40 per property.  |              |   |         | \$                    |              |
| <b>TOTAL FEES ENCLOSED=</b>  |              |   |         | <b>\$1,000.00</b>     |              |
|  |              |   |         | Amount to be refunded | \$           |
|  |              |   |         | Amount to be charged: | \$           |

a. ☐ No fee is believed to be due.

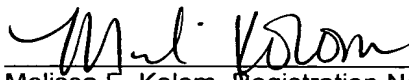
b. ☒ Please charge Deposit Account No. 12-1216 the total fee indicated above.

c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 12-1216. A duplicate copy of this sheet is enclosed.

**NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status.**

SEND ALL CORRESPONDENCE TO CUSTOMER NO. 23460:

**23460**

  
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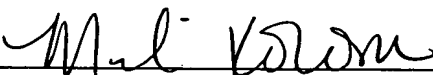
Date: May 26, 2006

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| U.S. APPLICATION NO. (if known, see 37 CFR 1.5)<br>Unassigned<br><div style="font-size: 24pt; font-weight: bold; margin-top: 10px;">10/580992</div> | INTERNATIONAL APPLICATION NO.<br>PCT/EP2004/013451 | ATTORNEY DOCKET NO.<br>252412 | CLIENT REF. NO.<br>PA36014US-<br>FZJH903ver |
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| EV709904617US<br><small>"Express Mail" Mailing Number</small> | May 26, 2006<br><small>Date of Deposit</small> |
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I hereby certify that this transmittal document with respect to the U.S. national phase of the above-referenced International Patent Application, including all of the items listed thereon as enclosures, is being deposited with the United States Postal Service "Express Mail Post Office to Addressee" Service under 37 CFR 1.10 on the date indicated above and is addressed to Mail Stop PCT, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

|   |  |
|---|--|
| Melissa E. Kolom  |  |
| <small>Typed or Printed Name of Person Making Deposit</small> | <small>Signature of Person Making Deposit</small>                                  |